

## PATENT COOPERATION TREATY

PCT

NOTIFICATION OF THE RECORDING  
OF A CHANGE(PCT Rule 92bis.1 and  
Administrative Instructions, Section 422)

From the INTERNATIONAL BUREAU

To:

MARTIN, Jean-Jacques  
Cabinet Regimbeau  
20, rue de Chazelles  
F-75847 Paris  
FRANCE

<b>Date of mailing</b> (day/month/year) 09 April 2001 (09.04.01)	<b>IMPORTANT NOTIFICATION</b>
<b>Applicant's or agent's file reference</b> 340289/18483	
<b>International application No.</b> PCT/IB99/01730	<b>International filing date</b> (day/month/year) 12 October 1999 (12.10.99)

## 1. The following indications appeared on record concerning:

☐ the applicant    ☐ the inventor    ☒ the agent    ☐ the common representative

<b>Name and Address</b> MARTIN, Jean-Jacques Cabinet Regimbeau 26, avenue Kléber F-75116 Paris France	<b>State of Nationality</b>	<b>State of Residence</b>
	<b>Telephone No.</b> 01-45-00-92-02	
	<b>Facsimile No.</b> 01-45-00-46-12	
	<b>Teleprinter No.</b>	

## 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

☐ the person    ☐ the name    ☒ the address    ☐ the nationality    ☐ the residence

<b>Name and Address</b> MARTIN, Jean-Jacques Cabinet Regimbeau 20, rue de Chazelles F-75847 Paris France	<b>State of Nationality</b>	<b>State of Residence</b>
	<b>Telephone No.</b> 33 1 44 29 35 00	
	<b>Facsimile No.</b> 33 1 44 29 35 99	
	<b>Teleprinter No.</b>	

## 3. Further observations, if necessary:

## 4. A copy of this notification has been sent to:

<input checked="" type="checkbox"/> the receiving Office	<input type="checkbox"/> the designated Offices concerned
<input type="checkbox"/> the International Searching Authority	<input checked="" type="checkbox"/> the elected Offices concerned
<input checked="" type="checkbox"/> the International Preliminary Examining Authority	<input type="checkbox"/> other:

<b>The International Bureau of WIPO</b> 34, chemin des Colombettes 1211 Geneva 20, Switzerland  Facsimile No.: (41-22) 740.14.35	<b>Authorized officer</b>  R. Chrem  Telephone No.: (41-22) 338.83.38
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## PATENT COOPERATION TREATY

PCT

## NOTIFICATION OF ELECTION

(PCT Rule 61.2)

From the INTERNATIONAL BUREAU

To:

Assistant Commissioner for Patents  
 United States Patent and Trademark  
 Office  
 Box PCT  
 Washington, D.C.20231  
 ETATS-UNIS D'AMERIQUE

in its capacity as elected Office

<b>Date of mailing</b> (day/month/year) 05 June 2000 (05.06.00)	
<b>International application No.</b> PCT/IB99/01730	<b>Applicant's or agent's file reference</b> 340289/18483
<b>International filing date</b> (day/month/year) 12 October 1999 (12.10.99)	<b>Priority date</b> (day/month/year) 13 October 1998 (13.10.98)
<b>Applicant</b> BLUMENFELD, Marta et al	

1. The designated Office is hereby notified of its election made:

☒ in the demand filed with the International Preliminary Examining Authority on:  
 08 May 2000 (08.05.00)

☐ in a notice effecting later election filed with the International Bureau on:

2. The election ☒ was  
☐ was not

made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).

<b>The International Bureau of WIPO</b> 34, chemin des Colombettes 1211 Geneva 20, Switzerland  Facsimile No.: (41-22) 740.14.35	Authorized officer  Olivia RANAIVOJAONA  Telephone No.: (41-22) 338.83.38
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## PATENT COOPERATION TREATY

PCT

From the INTERNATIONAL BUREAU

## NOTIFICATION RELATING TO PRIORITY CLAIM

(PCT Rules 26bis.1 and 26bis.2 and  
Administrative Instructions, Sections 402 and 409)

To:

MARTIN, Jean-Jacques  
Cabinet Regimbeau  
26, avenue Kléber  
F-75116 Paris  
FRANCE

Date of mailing (day/month/year) 17 December 1999 (17.12.99)	<b>IMPORTANT NOTIFICATION</b>
Applicant's or agent's file reference 340289/18483	
International application No. PCT/IB99/01730	International filing date (day/month/year) 12 October 1999 (12.10.99)
Applicant GENSET et al	

The applicant is hereby **notified** of the following in respect of the priority claim(s) made in the international application.

1. ☒ **Correction of priority claim.** In accordance with the applicant's notice received on: 10 December 1999 (10.12.99), the following priority claim has been corrected to read as follows:

US 13 October 1998 (13.10.98) 60/103,955

- ☐ even though the indication of the number of the earlier application is missing.  
☐ even though the following indication in the priority claim is not the same as the corresponding indication appearing in the priority document:

2. ☐ **Addition of priority claim.** In accordance with the applicant's notice received on: , the following priority claim has been added:

- ☐ even though the indication of the number of the earlier application is missing.  
☐ even though the following indication in the priority claim is not the same as the corresponding indication appearing in the priority document:

3. ☐ As a **result of the correction and/or addition** of (a) priority claim(s) under items 1 and/or 2, the (earliest) priority date is:

4. ☐ **Priority claim considered not to have been made.**

- ☐ The applicant failed to respond to the Invitation under Rule 26bis.2(a) (Form PCT/IB/316) within the prescribed time limit.  
☐ The applicant's notice was received after the expiration of the prescribed time limit under Rule 26bis.1(a).  
☐ The applicant's notice failed to correct the priority claim so as to comply with the requirements of Rule 4.10.

The applicant may, before the technical preparations for international publication have been completed and subject to the payment of a fee, request the International Bureau to publish, together with the international application, information concerning the priority claim. See Rule 26bis.2(c) and the PCT Applicant's Guide, Volume I, Annex B2(1B).

5. ☒ In case where **multiple priorities** have been claimed, the above item(s) relate to the following priority claim(s):

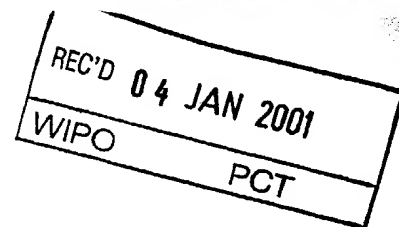
US 13 October 1998 (13.10.98) 60/103,955

6. A copy of this notification has been sent to the receiving Office and

- ☒ to the International Searching Authority (where the international search report has not yet been issued).  
☒ the designated Offices (which have already been notified of the receipt of the record copy).

<b>The International Bureau of WIPO</b> 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Authorized officer  R. Chrem
Facsimile No. (41-22) 740.14.35	Telephone No. (41-22) 338.83.38

# TENT COOPERATION TREATY PCT



## INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

Applicant's or agent's file reference <b>340289/18483</b>	<b>FOR FURTHER ACTION</b> See Notification of Transmittal of International Preliminary Examination Report (Form PCT/IPEA/416)	
International application No. <b>PCT/IB99/01730</b>	International filing date (day/month/year) <b>12/10/1999</b>	Priority date (day/month/year) <b>13/10/1998</b>
International Patent Classification (IPC) or national classification and IPC <b>C12N15/12</b>		
Applicant <b>GENSET et al.</b>		

1. This international preliminary examination report has been prepared by this International Preliminary Examining Authority and is transmitted to the applicant according to Article 36.
2. This REPORT consists of a total of 7 sheets, including this cover sheet.
 

☐ This report is also accompanied by ANNEXES, i.e. sheets of the description, claims and/or drawings which have been amended and are the basis for this report and/or sheets containing rectifications made before this Authority (see Rule 70.16 and Section 607 of the Administrative Instructions under the PCT).

These annexes consist of a total of    sheets.

3. This report contains indications relating to the following items:
 

I    ☒ Basis of the report

II   ☐ Priority

III ☒ Non-establishment of opinion with regard to novelty, inventive step and industrial applicability

IV   ☐ Lack of unity of invention

V    ☒ Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

VI   ☐ Certain documents cited

VII ☐ Certain defects in the international application

VIII ☒ Certain observations on the international application

Date of submission of the demand  <b>08/05/2000</b>	Date of completion of this report  <b>28.12.2000</b>
Name and mailing address of the international preliminary examining authority: European Patent Office D-80298 Munich Tel. +49 89 2399 - 0 Tx: 523656 epmu d Fax: +49 89 2399 - 4465	Authorized officer  <b>Bretherick, J</b>  Telephone No. +49 89 2399 8415



**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT**

International application No. PCT/IB99/01730

**I. Basis of the report**

1. This report has been drawn on the basis of *(substitute sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to the report since they do not contain amendments (Rules 70.16 and 70.17).):*

**Description, pages:**

1-186 as originally filed

**Claims, No.:**

1-57 as originally filed

**Drawings, sheets:**

1/12-12/12 as originally filed

**Sequence listing part of the description, pages:**

1-62, as originally filed

2. With regard to the **language**, all the elements marked above were available or furnished to this Authority in the language in which the international application was filed, unless otherwise indicated under this item.

These elements were available or furnished to this Authority in the following language: , which is:

- ☐ the language of a translation furnished for the purposes of the international search (under Rule 23.1(b)).  
☐ the language of publication of the international application (under Rule 48.3(b)).  
☐ the language of a translation furnished for the purposes of international preliminary examination (under Rule 55.2 and/or 55.3).

3. With regard to any **nucleotide and/or amino acid sequence** disclosed in the international application, the international preliminary examination was carried out on the basis of the sequence listing:

- ☒ contained in the international application in written form.  
☐ filed together with the international application in computer readable form.  
☐ furnished subsequently to this Authority in written form.  
☒ furnished subsequently to this Authority in computer readable form.  
☒ The statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.  
☐ The statement that the information recorded in computer readable form is identical to the written sequence listing has been furnished.

4. The amendments have resulted in the cancellation of:

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT**

International application No. PCT/IB99/01730

- ☐ the description,      pages:  
☐ the claims,      Nos.:  
☐ the drawings,      sheets:

5. ☐ This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed (Rule 70.2(c)):

*(Any replacement sheet containing such amendments must be referred to under item 1 and annexed to this report.)*

6. Additional observations, if necessary:

**III. Non-establishment of opinion with regard to novelty, inventive step and industrial applicability**

1. The questions whether the claimed invention appears to be novel, to involve an inventive step (to be non-obvious), or to be industrially applicable have not been examined in respect of:

- ☐ the entire international application.  
☒ claims Nos. 2,5,48,49, completely, 1,3,4,6-25,28-47,50-57, partially.

because:

- ☐ the said international application, or the said claims Nos. relate to the following subject matter which does not require an international preliminary examination (*specify*):
- ☐ the description, claims or drawings (*indicate particular elements below*) or said claims Nos. are so unclear that no meaningful opinion could be formed (*specify*):
- ☐ the claims, or said claims Nos. are so inadequately supported by the description that no meaningful opinion could be formed.
- ☒ no international search report has been established for the said claims Nos. As above..
2. A meaningful international preliminary examination report cannot be carried out due to the failure of the nucleotide and/or amino acid sequence listing to comply with the standard provided for in Annex C of the Administrative Instructions:
- ☐ the written form has not been furnished or does not comply with the standard.
- ☐ the computer readable form has not been furnished or does not comply with the standard.

**V. Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement**

# INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/IB99/01730

## 1. Statement

Novelty (N)	Yes:	Claims	3,4,6-14,16-22,25-27,28-44,46,47,50,57.
	No:	Claims	1,15,23,24,45,51-56
Inventive step (IS)	Yes:	Claims	
	No:	Claims	1,3,4,6-25,28-47,50-57.
Industrial applicability (IA)	Yes:	Claims	1,3,4,6-14,16-47,50-57
	No:	Claims	

## 2. Citations and explanations **see separate sheet**

## VIII. Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:  
**see separate sheet**

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET**

International application No. PCT/IB99/01730

1. **Regarding Part I**, the application contains a 62-page sequence listing in addition to the description, claims and figures.
2. **Regarding Part III**, the claims searched have been examined only with respect to the first invention mentioned in Box C of the International Search Report. There is also a further examination restriction with respect to claims 16-18, since the subject-matter was found to be unsearchable in its broadest sense.
3. **Regarding Part VIII**, Art. 6 PCT:
  - a. Claim 1 is not clear, since the selection of at least 12 nucleotides from the sequences at the individual positions mentioned or their complements results in a huge number of possibilities. This also applies to claims 26 and 27, 51 and 57 and dependencies, since these also refers to such subject-matter.
  - b. The same applies **mutatis mutandis** to claim 3, defining a contiguous span of 8-50 nucleotides of anyone of SEQ ID NOS 1-3 or their complements, which are only further defined by reference to including a G713 or q31-q33-related biallelic marker in said sequence without defining what these are in structural terms within the claims. Claims 4, 6, 7-14, 16-25, 28-44, 47, 50, 52 and dependencies are also not clear for the same reasons since they either refer to claim 3 or are not defined in terms anything other than by reference to arbitrary attributes.
  - c. Claim 15 is unclear, since it refers to polynucleotides encoding a polypeptide comprising a contiguous span of at least 6 amino acids of SEQ ID NO:5, without either defining the polypeptide **per se**, or its function.  
This also applies with respect to claim 57, part (f).
  - d. Since claims 16-18 refer to an almost limitless number of possible combinations which have not been searched, they have been interpreted as being limited to the subject-matter of Table 2 on page 148 of the application. Even so, the claims are not clearly defined since the polynucleotide of claim 16 is defined solely in terms of its suitability for use in a genotyping assay for determining the identity of the nucleotide at a G713- or 13q31-q33-related biallelic marker, or the complement thereof. Claims 17 and 18 do not add any further clarifying matter.



**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET**

International application No. PCT/IB99/01730

4. The above notwithstanding, **Regarding Part V, Art. 33 PCT:**
- a. In as far as claims 51-56 are concerned with computer readable media, the subject-matter thereof is considered to be distinguished from the art merely by the non-technical feature of containing stored information. This has been treated as subject-matter in respect of which an international preliminary examination Report is not mandatory, Art. 34(4)(a)(i), R. 67(v) PCT. Nevertheless, an opinion has been given on this subject-matter. It should be born in mind that such subject-matter may not be patentable before certain regional authorities under the Chapter II PCT system.
  - b. The subject-matter of claim 1 is not new under Art. 33(1)(2) PCT over the disclosure of Li et al. (1993) Genomics Vol. 16 pp. 572-579, and the associated database EMBL/GENBANK Accession No. L10374 Sequence reference HSMRNAC. This sequence has 99.756% identity with SEQ. ID. NO: 4. This also applies with respect to the subject-matter of claims 15, 23, 24, 45, since these are the corresponding amino acid sequences which are comprised in this cDNA.

The subject-matter of claims 51-56 is also not new, since the insertion of said art sequence into the database involves its storage on a computer, with the capacity to carry out sequence comparisons with others and to observe polymorphisms and other sequence differences.

Claim 57 is considered not to involve an inventive step under Art. 33(1)(3) PCT. The sequence claimed is compared to a generic reference sequence in such a system. Whilst it is not documented whether the art sequence has been used in this way previously, its use would be a clear possibility for the person skilled in the art for any application.

- c. The remaining claimed subject-matter is not considered to involve an inventive step under Art. 33(1)(3) PCT.

In the case of generic and specific nucleotide sequences defined for example, in terms of arbitrary sequence length in a defined SEQ ID NO and further defined according to having G713- or 13q31-q33-related biallelic markers in such a

**INTERNATIONAL PRELIMINARY  
EXAMINATION REPORT - SEPARATE SHEET**

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International application No. PCT/IB99/01730

sequence, the use of biallelic markers is considered to be useful in linkage studies, (Kruglyak (1997), nature genetics Vol. 17 pp. 21-24). There is also suggestive evidence in the art for the linkage of schizophrenia to markers on chromosome 13, (13q14.1-q32) (Lin et al. (1997) Human Genetics Vol. 99 pp. 417-420). An area of potential linkage on chromosome region 13q32 to schizophrenia is also disclosed in Detera-Wadleigh et al. (1999) PNAS USA Vol. 96 pp. 5604-5609. The current broadly and specifically formulated claims are not considered to involve an inventive step, since the skilled person would be firstly prompted to look in the region of 13q32 for such markers and secondly, even in the case where specific markers are claimed, there is no surprising effect associated with the specific sequence isolated using standard art techniques.

This position applies mutatis mutandis to the subject-matter involving the use of such markers in for example, methods involving linkage analysis and antibodies etc. directed to specific or generic amino acid sequences defined by coding regions thereby defined.

**FOR THE PURPOSES OF INFORMATION ONLY**

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
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DK	Denmark	LR	Liberia	SG	Singapore		
EE	Estonia						

# INTERNATIONAL SEARCH REPORT

International Application No  
PCT/IB 99/01730

## A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 C12N15/12 C12N5/10 C12N1/21 A01K67/027 C12N15/00  
C12Q1/68 C07K16/18

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C07K C12Q

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	LI S -H ET AL: "NOVEL TRIPLET REPEAT CONTAINING GENES IN HUMAN BRAIN: CLONING, EXPRESSION, AND LENGTH POLYMORPHISMS" GENOMICS, US, ACADEMIC PRESS, SAN DIEGO, vol. 16, no. 3, 1 June 1993 (1993-06-01), pages 572-579, XP000196589 ISSN: 0888-7543 cited in the application	1,15,45, 46, 50-54, 56,57
Y	the whole document  -& EMBL/GENBANK DATABASES Accession no L10374 Sequence reference HSMRNAC 23 June 1993 XP002134451  --- -/--	3,4, 6-14, 16-44, 47,55

☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

### \* Special categories of cited documents:

"A" document defining the general state of the art which is not considered to be of particular relevance

"E" earlier document but published on or after the international filing date

"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.

"Z" document member of the same patent family

Date of the actual completion of the international search

10 April 2000

Date of mailing of the international search report

10. 07. 00

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2  
NL - 2280 HV Rijswijk  
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  
Fax: (+31-70) 340-3016

Authorized officer

VAN DER SCHAAL C.A.

# INTERNATIONAL SEARCH REPORT

International Application No.

PCT/IB 99/01730

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	<p>WANG D G ET AL: "Large-scale identification, mapping, and genotyping of single-nucleotide polymorphisms in the human genome"</p> <p>SCIENCE,US,AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE,, vol. 280, 1 January 1998 (1998-01-01), pages 1077-1082, XP002089398 ISSN: 0036-8075 the whole document</p> <p style="text-align: center;">---</p>	<p>3,4, 6-14, 16-44, 47,55</p>
Y	<p>KRUGLYAK L: "PASSAGE TEXT. THE USE OF A GENETIC MAP OF BIALLELIC MARKERS IN LINKAGE STUDIES"</p> <p>NATURE GENETICS,US,NEW YORK, NY, vol. 17, no. 1, 1 September 1997 (1997-09-01), pages 22-24, XP002050647 ISSN: 1061-4036 the whole document</p> <p style="text-align: center;">---</p>	<p>3,4, 6-14, 16-25, 28-44, 47,55</p>
Y	<p>WO 98 11222 A (SINAI SCHOOL MEDICINE) 19 March 1998 (1998-03-19) page 17, paragraph 1 page 23, paragraph 3</p> <p style="text-align: center;">---</p>	<p>26,27</p>
Y	<p>DETERA-WADLEIGH S D ET AL: "A high-density genome scan detects evidence for a bipolar-disorder susceptibility locus on 13q32 and other potential loci on 1q32 and 18p11.2."</p> <p>PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES OF AMERICA, (1999 MAY 11) 96 (10) 5604-9., XP002134450 the whole document</p> <p style="text-align: center;">---</p>	<p>42,47</p>
Y	<p>MING-WEI LIN ET AL: "SUGGESTIVE EVIDENCE FOR LINKAGE OF SCHIZOPHRENIA TO MARKERS ON CHROMOSOME 13 IN CAUCASIAN BUT NOT ORIENTAL POPULATIONS"</p> <p>HUMAN GENETICS,DE,BERLIN, vol. 99, no. 3, March 1997 (1997-03), pages 417-420, XP000876937 the whole document</p> <p style="text-align: center;">---</p>	<p>42,47</p>
X	<p>EMBL/GENBANK DATABASES Accession no AA989298 Sequence ID AA989298 27 August 1998 "National Cancer INstitute, Cancer Genome Anatomy Project..." XP002135005 the whole document</p> <p style="text-align: center;">---</p>	<p>1,45,46</p>
	-/--	

# INTERNATIONAL SEARCH REPORT

International Application No  
PCT/IB 99/01730

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>EMBL DATABASE Accession no B75361 Sequence ID B75361 14 Januari 1998 ADAMS M ET AL: "Use of BAC end sequences for sequence-ready map building" XP002135179 the whole document -----</p>	1

# INTERNATIONAL SEARCH REPORT

national application No.  
PCT/IB 99/01730

## Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:  
Although claims 51-56 could be considered as a mere presentation of information or a computer program, Rule 39.1 (v and vi) PCT, the search has been carried out as far as possible in our systematic documentation.
2. ☒ Claims Nos.:  
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:  
see FURTHER INFORMATION sheet PCT/ISA/210
3. ☐ Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

*see additional sheet*

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☒ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  
1, 3, 4, 6-25, 28-47, 50-57 partially, 26, 27 completely

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.  
☐ No protest accompanied the payment of additional search fees.

## INTERNATIONAL SEARCH REPORT

International Application No. PCT/IB 99/01730

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

1. Claims: 1 3 4 6-25 28-47 50-57 partially 26 27 completely

human G713, (fragments of) nucleotides encoding G713  
including biallelic markers derived from these sequences,  
antibodies against G713 and their uses

2. Claims: 1 51 53-57 partially 2 completely

(fragments of) nucleotide sequence SEQ ID NO 31 and its use

3. Claims: 1 3 4 6-14 16-25 28-44 47 50-  
57 partially 5 48 49 completely

(fragments of) 13q31-q33 biallelic marker containing  
sequences SEQ ID NO 32-69 and their uses

4. Claims: 1 15 45 46 51-57 partially

murine homologue of G713, nucleotides encoding it,  
antibodies against it and their uses



## INTERNATIONAL SEARCH REPORT

International Application No. PCT/IB 99/01730

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Present claims 16-18 relate to an extremely large number of possible compoundss. In fact, the claims contain so many options, that a lack of clarity (and/or conciseness) within the meaning of Article 6 PCT arises to such an extent as to render a meaningful search of the claims impossible. Consequently, the search has been carried out for those parts of the application which do appear to be clear (and/or concise), namely the polynucleotides mentioned in tabel 2

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

Orma in patent family members

PCT/IB 99/01730

Form PCT/ISA/210 (patent family annex) (July 1992)

PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference <b>340289/18483</b>	<b>FOR FURTHER ACTION</b> see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.	
International application No. <b>PCT/ IB 99/ 01730</b>	International filing date (day/month/year) <b>12/10/1999</b>	(Earliest) Priority Date (day/month/year) <b>13/10/1998</b>
Applicant <b>GENSET et al.</b>		

This International Search Report has been prepared by this International Searching Authority and is transmitted to the applicant according to Article 18. A copy is being transmitted to the International Bureau.

This International Search Report consists of a total of 7 sheets.  
☐ It is also accompanied by a copy of each prior art document cited in this report.

1. Basis of the report

a. With regard to the **language**, the international search was carried out on the basis of the international application in the language in which it was filed, unless otherwise indicated under this item.

☐ the international search was carried out on the basis of a translation of the international application furnished to this Authority (Rule 23.1(b)).

b. With regard to any **nucleotide and/or amino acid sequence** disclosed in the international application, the international search was carried out on the basis of the sequence listing :

☒ contained in the international application in written form.

☐ filed together with the international application in computer readable form.

☐ furnished subsequently to this Authority in written form.

☒ furnished subsequently to this Authority in computer readable form.

☒ the statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.

☐ the statement that the information recorded in computer readable form is identical to the written sequence listing has been furnished

2. ☒ **Certain claims were found unsearchable** (See Box I).

3. ☒ **Unity of invention is lacking** (see Box II)

4. With regard to the **title**,

☒ the text is approved as submitted by the applicant.

☐ the text has been established by this Authority to read as follows:

5. With regard to the **abstract**,

☒ the text is approved as submitted by the applicant.

☐ the text has been established, according to Rule 38.2(b), by this Authority as it appears in Box III. The applicant may, within one month from the date of mailing of this international search report, submit comments to this Authority.

6. The figure of the **drawings** to be published with the abstract is Figure No.

☐ as suggested by the applicant.

☐ because the applicant failed to suggest a figure.

☐ because this figure better characterizes the invention.

☒ None of the figures.

**Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)**

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:  
Although claims 51-56 could be considered as a mere presentation of information or a computer program, Rule 39.1 (v and vi) PCT, the search has been carried out as far as possible in our systematic documentation.
2. ☒ Claims Nos.:  
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:  
see FURTHER INFORMATION sheet PCT/ISA/210
3. ☐ Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

**Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)**

This International Searching Authority found multiple inventions in this international application, as follows:

see additional sheet

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☒ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

1, 3, 4, 6-25, 28-47, 50-57 partially, 26, 27 completely

**Remark on Protest**

- ☐ The additional search fees were accompanied by the applicant's protest.  
☐ No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

1. Claims: 1 3 4 6-25 28-47 50-57 partially 26 27 completely  
human G713, (fragments of) nucleotides encoding G713  
including biallelic markers derived from these sequences,  
antibodies against G713 and their uses
2. Claims: 1 51 53-57 partially 2 completely  
(fragments of) nucleotide sequence SEQ ID NO 31 and its use
3. Claims: 1 3 4 6-14 16-25 28-44 47 50-  
57 partially 5 48 49 completely  
  
(fragments of) 13q31-q33 biallelic marker containing  
sequences SEQ ID NO 32-69 and their uses
4. Claims: 1 15 45 46 51-57 partially  
murine homologue of G713, nucleotides encoding it,  
antibodies against it and their uses

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

## Continuation of Box I.2

Present claims 16-18 relate to an extremely large number of possible compoundss. In fact, the claims contain so many options, that a lack of clarity (and/or conciseness) within the meaning of Article 6 PCT arises to such an extent as to render a meaningful search of the claims impossible. Consequently, the search has been carried out for those parts of the application which do appear to be clear (and/or concise), namely the polynucleotides mentioned in tabel 2

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

## INTERNATIONAL SEARCH REPORT

International Application No

PC 99/01730

## A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 C12N15/12 C12N5/10 C12N1/21 A01K67/027 C12N15/00  
 C12Q1/68 C07K16/18

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C07K C12Q

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	LI S -H ET AL: "NOVEL TRIPLET REPEAT CONTAINING GENES IN HUMAN BRAIN: CLONING, EXPRESSION, AND LENGTH POLYMORPHISMS" GENOMICS, US, ACADEMIC PRESS, SAN DIEGO, vol. 16, no. 3, 1 June 1993 (1993-06-01), pages 572-579, XP000196589 ISSN: 0888-7543 cited in the application	1, 15, 45, 46, 50-54, 56, 57
Y	the whole document  -& EMBL/GENBANK DATABASES Accession no L10374 Sequence reference HSMRNAC 23 June 1993 XP002134451  --- -/-	3, 4, 6-14, 16-44, 47, 55

☒ Further documents are listed in the continuation of box C.☒ Patent family members are listed in annex.

## \* Special categories of cited documents :

- "A" document defining the general state of the art which is not considered to be of particular relevance  
 "E" earlier document but published on or after the international filing date  
 "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)  
 "O" document referring to an oral disclosure, use, exhibition or other means  
 "P" document published prior to the international filing date but later than the priority date claimed

- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention  
 "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone  
 "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.  
 "&" document member of the same patent family

Date of the actual completion of the international search

10 April 2000

Date of mailing of the international search report

10.07.00

Name and mailing address of the ISA

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 NL - 2280 HV Rijswijk  
 Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  
 Fax: (+31-70) 340-3016

Authorized officer

VAN DER SCHAAL C.A.

## INTERNATIONAL SEARCH REPORT

International Application No

PC 99/01730

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	<p>WANG D G ET AL: "Large-scale identification, mapping, and genotyping of single-nucleotide polymorphisms in the human genome"</p> <p>SCIENCE,US,AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE,, vol. 280, 1 January 1998 (1998-01-01), pages 1077-1082, XP002089398 ISSN: 0036-8075 the whole document</p>	3,4, 6-14, 16-44, 47,55
Y	<p>--- KRUGLYAK L: "PASSAGE TEXT. THE USE OF A GENETIC MAP OF BIALLELIC MARKERS IN LINKAGE STUDIES"</p> <p>NATURE GENETICS,US,NEW YORK, NY, vol. 17, no. 1, 1 September 1997 (1997-09-01), pages 22-24, XP002050647 ISSN: 1061-4036 the whole document</p>	3,4, 6-14, 16-25, 28-44, 47,55
Y	<p>--- WO 98 11222 A (SINAI SCHOOL MEDICINE) 19 March 1998 (1998-03-19) page 17, paragraph 1 page 23, paragraph 3</p>	26,27
Y	<p>--- DETERA-WADLEIGH S D ET AL: "A high-density genome scan detects evidence for a bipolar-disorder susceptibility locus on 13q32 and other potential loci on 1q32 and 18p11.2."</p> <p>PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES OF AMERICA, (1999 MAY 11) 96 (10) 5604-9., XP002134450 the whole document</p>	42,47
Y	<p>--- MING-WEI LIN ET AL: "SUGGESTIVE EVIDENCE FOR LINKAGE OF SCHIZOPHRENIA TO MARKERS ON CHROMOSOME 13 IN CAUCASIAN BUT NOT ORIENTAL POPULATIONS"</p> <p>HUMAN GENETICS,DE,BERLIN, vol. 99, no. 3, March 1997 (1997-03), pages 417-420, XP000876937 the whole document</p>	42,47
X	<p>--- EMBL/GENBANK DATABASES Accession no AA989298 Sequence ID AA989298 27 August 1998 "National Cancer INstitute, Cancer Genome Anatomy Project..." XP002135005 the whole document</p> <p>--- -/--</p>	1,45,46



# INTERNATIONAL SEARCH REPORT

International Application No

PCT/B 99/01730

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>EMBL DATABASE Accession no B75361 Sequence  ID B75361  14 Januari 1998  ADAMS M ET AL: "Use of BAC end sequences  for sequence-ready map building"  XP002135179  the whole document  -----</p>	1

### Information on patent family members

PC B 99/01730

Patent document  
cited in search report

Publication  
date

Patent family member(s)

Publication date

WO 9811222

A

19-03-1998

AU

4480997 A

02-04-1998

# PCT

## INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference <b>340289/18483</b>	<b>FOR FURTHER ACTION</b> see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.	
International application No. <b>PCT/IB 99/ 01730</b>	International filing date (day/month/year) <b>12/10/1999</b>	(Earliest) Priority Date (day/month/year) <b>13/10/1998</b>
Applicant <b>GENSET et al.</b>		

This International Search Report has been prepared by this International Searching Authority and is transmitted to the applicant according to Article 18. A copy is being transmitted to the International Bureau.

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☒ the statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.

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3. ☒ **Unity of invention is lacking** (see Box II).

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6. The figure of the **drawings** to be published with the abstract is Figure No.

☐ as suggested by the applicant.

☐ because the applicant failed to suggest a figure.

☐ because this figure better characterizes the invention.

☒ None of the figures.

# INTERNATIONAL SEARCH REPORT

International application No.  
PCT/IB 99/01730

## Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:  
  
Although claims 51-56 could be considered as a mere presentation of information or a computer program, Rule 39.1 (v and vi) PCT, the search has been carried out as far as possible in our systematic documentation.
2. ☒ Claims Nos.:  
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:  
  
see FURTHER INFORMATION sheet PCT/ISA/210
3. ☐ Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

see additional sheet

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☒ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  
  
1, 3, 4, 6-25, 28-47, 50-57 partially, 26, 27 completely

### Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☐ No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

1. Claims: 1 3 4 6-25 28-47 50-57 partially 26 27 completely

human G713, (fragments of) nucleotides encoding G713  
including biallelic markers derived from these sequences,  
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## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

## Continuation of Box I.2

Present claims 16-18 relate to an extremely large number of possible compoundss. In fact, the claims contain so many options, that a lack of clarity (and/or conciseness) within the meaning of Article 6 PCT arises to such an extent as to render a meaningful search of the claims impossible. Consequently, the search has been carried out for those parts of the application which do appear to be clear (and/or concise), namely the polynucleotides mentioned in tabel 2

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

## INTERNATIONAL SEARCH REPORT

 International Application No  
 PC1,1B 99/01730

## A. CLASSIFICATION OF SUBJECT MATTER

 IPC 7 C12N15/12 C12N5/10 C12N1/21 A01K67/027 C12N15/00  
 C12Q1/68 C07K16/18

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C07K C12Q

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	LI S -H ET AL: "NOVEL TRIPLET REPEAT CONTAINING GENES IN HUMAN BRAIN: CLONING, EXPRESSION, AND LENGTH POLYMORPHISMS" GENOMICS,US,ACADEMIC PRESS, SAN DIEGO, vol. 16, no. 3, 1 June 1993 (1993-06-01), pages 572-579, XP000196589 ISSN: 0888-7543 cited in the application	1,15,45, 46, 50-54, 56,57
Y	the whole document  -& EMBL/GENBANK DATABASES Accession no L10374 Sequence reference HSMRNAC 23 June 1993 XP002134451  -/-	3,4, 6-14, 16-44, 47,55

☒ Further documents are listed in the continuation of box C.☒ Patent family members are listed in annex.

## \* Special categories of cited documents:

- \*A\* document defining the general state of the art which is not considered to be of particular relevance
- \*E\* earlier document but published on or after the international filing date
- \*L\* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- \*O\* document referring to an oral disclosure, use, exhibition or other means
- \*P\* document published prior to the international filing date but later than the priority date claimed

- \*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- \*X\* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- \*Y\* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- \*Z\* document member of the same patent family

Date of the actual completion of the international search

10 April 2000

Date of mailing of the international search report

10.07.00

Name and mailing address of the ISA

 European Patent Office, P.B. 5818 Patentlaan 2  
 NL - 2280 HV Rijswijk  
 Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  
 Fax: (+31-70) 340-3016

Authorized officer

VAN DER SCHAAL C.A.

## INTERNATIONAL SEARCH REPORT

 International Application No  
 PCT/IB 99/01730

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	<p>WANG D G ET AL: "Large-scale identification, mapping, and genotyping of single-nucleotide polymorphisms in the human genome"</p> <p>SCIENCE,US,AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE,, vol. 280, 1 January 1998 (1998-01-01), pages 1077-1082, XP002089398 ISSN: 0036-8075 the whole document</p> <p>---</p>	<p>3,4, 6-14, 16-44, 47,55</p>
Y	<p>KRUGLYAK L: "PASSAGE TEXT. THE USE OF A GENETIC MAP OF BIALLELIC MARKERS IN LINKAGE STUDIES"</p> <p>NATURE GENETICS,US,NEW YORK, NY, vol. 17, no. 1, 1 September 1997 (1997-09-01), pages 22-24, XP002050647 ISSN: 1061-4036 the whole document</p> <p>---</p>	<p>3,4, 6-14, 16-25, 28-44, 47,55</p>
Y	<p>WO 98 11222 A (SINAI SCHOOL MEDICINE) 19 March 1998 (1998-03-19) page 17, paragraph 1 page 23, paragraph 3</p> <p>---</p>	<p>26,27</p>
Y	<p>DETERA-WADLEIGH S D ET AL: "A high-density genome scan detects evidence for a bipolar-disorder susceptibility locus on 13q32 and other potential loci on 1q32 and 18p11.2."</p> <p>PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES OF AMERICA, (1999 MAY 11) 96 (10) 5604-9., XP002134450 the whole document</p> <p>---</p>	<p>42,47</p>
Y	<p>MING-WEI LIN ET AL: "SUGGESTIVE EVIDENCE FOR LINKAGE OF SCHIZOPHRENIA TO MARKERS ON CHROMOSOME 13 IN CAUCASIAN BUT NOT ORIENTAL POPULATIONS"</p> <p>HUMAN GENETICS,DE,BERLIN, vol. 99, no. 3, March 1997 (1997-03), pages 417-420, XP000876937 the whole document</p> <p>---</p>	<p>42,47</p>
X	<p>EMBL/GENBANK DATABASES Accession no AA989298 Sequence ID AA989298 27 August 1998 "National Cancer Institute, Cancer Genome Anatomy Project..." XP002135005 the whole document</p> <p>---</p>	<p>1,45,46</p>
	<p>---</p> <p>-/--</p>	



# INTERNATIONAL SEARCH REPORT

International Application No  
PCT/IB 99/01730

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>EMBL DATABASE Accession no B75361 Sequence ID B75361 14 Januari 1998 ADAMS M ET AL: "Use of BAC end sequences for sequence-ready map building" XP002135179 the whole document -----</p>	1

Inform on patent family members

PC, IB 99/01730

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9811222 A	19-03-1998	AU 4480997 A	02-04-1998
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